

FIG. 1A

KCNQ5 cDNA coding sequence

atgaaggatg tggagtcggg ccggggcagg gtgctgctga 40
actcggcagc cgccagggggc gacggcctgc tactgctggg 80
caccgcgcgc gccacgcttg gtggcggcgg cggtggcctg 120
agggagagcc gccgggggcaa gcaggggggcc cggatgagcc 160
tgctggggaa gccgctctct tacacgagta gccagagctg 200
ccggcgcaac gtcaagtacc ggcggggtgca gaactacctg 240
tacaacgtgc tggagagacc ccgcggctgg gcgttcctct 280
accacgcttt cgtttttctc cttgtctttg gttgcttgat 320
tttgtcagtg ttttctacca tccctgagca cacaaaattg 360
gcctcaagtt gcctcttgat cctggagttc gtgatgattg 400
tcgtctttgg tttggagttc atcattcgaa tctggtctgc 440
gggttgctgt tgtcgatata gaggatggca aggaagactg 480
aggtttgctc gaaagccctt ctgtgttata gataccattg 520
ttcttategc ttcaatagca gttgtttctg caaaaactca 560
gggtaatat tttgccacgt ctgcactcag aagtctccgt 600
ttcctacaga tcctccgcac ggtgcgcacg gaccgaaggg 640
gaggcacttg gaaattactg ggttcagtgg tttatgctca 680
cagcaaggaa ttaatcacag cttggtacat aggatttttg 720
gttcttattt tttcgtcttt ccttgtctat ctggtggaaa 760

FIG. 1B

aggatgccaa taaagagttt tctacatatg cagatgctct 800
ctggtggggc acaattacat tgacaactat tggctatgga 840
gacaaaactc ccctaacttg gctgggaaga ttgctttctg 880
caggctttgc actccttggc atttctttct ttgcacttcc 920
tgccggcatt cttggctcag gttttgcatt aaaagtacaa 960
gaacaacacc gccagaaaca ctttgagaaa agaaggaacc 1000
cagctgccaa cctcattcag tgtgtttggc gtagttacgc 1040
agctgatgag aaatctgttt ccattgcaac ctggaagcca 1080
cacttgaagg ccttgcacac ctgcagccct accaagaaag 1120
aacaagggga agcatcaagc agtcagaagc taagttttaa 1160
ggagcgagtg cgcattggcta gcccagggg ccagagtatt 1200
aagagccgac aagcctcagt aggtgacagg aggtccccaa 1240
gcaccgacat cacagccgag ggagtcacca ccaaagtgca 1280
gaagagctgg agcttcaacg accgaaccg cttccggccc 1320
tcgctgcgcc tcaaaagttc tcagccaaaa ccagtgatag 1360
atgctgacac agcccttggc actgatgatg tatatgatga 1400
aaaaggatgc cagtgtgatg tatcagtgga agacctcacc 1440
ccaccactta aaactgtcat tcgagctatc agaattatga 1480
aatttcatgt tgcaaaacgg aagtttaagg aaacgttacg 1520
tccatatgat gtaaaagatg tcattgaaca atattctgct 1560

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FIG. 1C

ggatcatctgg acatgtttgtg tagaattaaa agccttcaaa 1600
cacgtgttga tcaaattctt ggaaaagggc aaatcacatc 1640
agataagaag agccgagaga aaataacagc agaacatgag 1680
accacagacg atctcagtat gctcggtcgg gtggtcaagg 1720
ttgaaaaaca ggtacagtcc atagagtcca agctggactg 1760
cctactagac atctatcaac aggtccttcg gaaaggctct 1800
gcctcagccc tcgctttggc ttcattccag atcccacctt 1840
ttgaatgtga acagacatct gactatcaaa gcctgtgga 1880
tagcaaagat ctttcgggtt ccgcacaaaa cagtggctgc 1920
ttatccagat caactagtgc caacatctcg agaggcctgc 1960
agttcattct gacgccaaat gagttcagtg cccagacttt 2000
ctacgcgctt agccctacta tgcacagtca agcaacacag 2040
gtgccaatta gtcaaagcga tggctcagca gtggcagcca 2080
ccaacaccat tgcaaacc aaataacagg caccacagcc 2120
agcagcccca acaactttac agatcccacc tcctctccca 2160
gccatcaagc atctgcccag gccagaaact ctgcacccta 2200
accctgcagg cttacaggaa agcattttctg acgtcaccac 2240
ctgccttggt gcctccaagg aaaatgttca ggttgcacag 2280
tcaaattctca ccaaggaccg ttctatgagg aaaagctttg 2320
acatgggagg agaaactctg ttgtctgtct gtcccatggg 2360
gccgaaggac ttgggcaa atcttgtctgt gcaaaacctg 2400
atcaggtcga ccgaggaact gaatatataa ctttcaggga 2440

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FIG. 1D

gtgagtcaag tggctccaga ggcagccaag atttttacct 2480
caaatggagg gaatccaaat tgtttataac tgatgaagag 2520
gtgggtcccg aagagacaga gacagacact tttgatgccg 2560
caccgcagcc tgccagggaa gctgcctttg catcagactc 2600
tctaaggact ggaagggtcac gatcatctca gagcatttgt 2640
aaggcaggag aaagtacaga tgccctcagc ttgcctcatg 2680
tcaaactgaa ataa 2694

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FOH250" 02099860

FIG. 2A

KCNQ5 Protein Sequence

MKDVESGRGR	VLLNSAAARG	DGLLLLGT	ATLGGGGGGL	40
RESRRGKQGA	RMSLLGKPLS	YTSSQSCRN	VKYRRVQNYL	80
YNVLERPRGW	AFIYHAFVFL	LVFGCLILSV	FSTIPEHTKL	120
ASSCLLILEF	VMIVVFGLEF	IIRIWSAGCC	CRYRGWQGRL	160
RFARKPFCVI	DTIVLIASIA	VVSAKTQGNI	FATSALRSLR	200
FLQILRMVRM	DRRGGTWKLL	GSVVYAHSKE	LITAWYIGFL	240
VLIFSSFLVY	LVEKDANKEF	STYADALWWG	TITLTTIGYG	280
DKTPLTWLGR	LLSAGFALLG	ISFFALPAGI	LGSGFALKVQ	320
EQHRQKHFEK	RRNPAANLIQ	CVWRSYAADE	KSVSIATWKP	360
HLKALHTCSP	TKKEQGEASS	SQKLSFKERV	RMASPRGQSI	400
KSRQASVGDR	RSPSTDITAE	GSPTKVQKSW	SFNDRTRFRP	440
SLRLKSSQPK	PVIDADTALG	TDDVYDEKGC	QCDVSVEDLT	480
PPLKTVIRAI	RIMKFHVAKR	KFKETLRPYD	VKDVIEQYSA	520
GHLDMLCRIK	SLQTRVDQIL	GKGQITSDDK	SREKITAEHE	560
TTDDL SMLGR	VVKVEKQVQS	IESKLDCLLD	IYQQVLRKGS	600
ASALALASFQ	IPPFECEQTS	DYQSPVDSKD	LSGSAQN SGC	640
LSRSTSANIS	RGLQFILTPN	EFSAQTFYAL	SPTMHSQATQ	680
VPISQSDGSA	VAATNTIANQ	INTAPKPAAP	TTLQIPPPLP	720
AIKHLPRPET	LHPNPAGLQE	SISDVTTCCLV	ASKENVQVAQ	760
SNLTKDRSMR	KSFDMGGETL	LSVCPMVPKD	LGKSLSVQNL	800
IRSTEELNIQ	LSGSESSGSR	GSQDFYPKWR	ESKLFITDEE	840

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FIG. 2B

VGPEETETDT FDAAPQPARE AAFASDSLRT GRSRSSQSIC	880
KAGESTDALS LPHVKLK	897

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FIG. 3

Alternative Splice Exon 1

TGG	GGA	CAG	TGG	ACA	TTG	CGT
Trp	Gly	Gln	Trp	Thr	Leu	Arg

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FIG. 4A

3' UTR

gttcttcatt	ttctttccag	gcatagcagt	tcttttagcca	40
tacatatcat	tgcatgaact	atttcgaaag	cccttctaaa	80
aagttgaaat	tgcaagaatc	gggaagaaca	tgaaaggcag	120
tttataagcc	cgttaccttt	taattgcatg	aaaatgcatg	160
tttagggatg	gctaaaattc	caaggtgcat	cgacattaac	200
ccactcattt	agtaatgtac	cttgagttaa	aaagcctgag	240
aaaccaaaca	cagctaattgc	tatgggggtgt	atgaatatgt	280
caagtttagg	tcattttagaa	gatttgacac	tgtattttga	320
aattatgagt	aaacaccttc	aaatttcagg	catttctgct	360
ttgtgactaa	atacaaacta	cattttcaag	attaggccat	400
aatgtatatt	taaacacaat	ggctatcaac	agctgctaata	440
aaggatatcaa	ctaaagcaga	attgggggaat	aatagaaatg	480
gctgcttatt	tcaagatata	tttgccaacc	cattcctatt	520
cagtcatttt	attattaatg	taatttgaat	gtcaatttgt	560
gtgcttttgg	tgatttagcg	ctgtggcaag	caattttgca	600
catcattttc	atgttggtct	ttatgacaag	aatgtttctc	640
aattagaaaa	tgtgcaaata	atgaaattca	gggccagtga	680
ggcaaataga	ctatctgaca	tatttgactt	tatgaaaaca	720
tattgcctga	tggcagaatc	aactttataa	gtggtcaact	760
tctacacaag	cgtatgaaat	actggtcagt	agaacagcca	800

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FIG. 4B

ttgtgattgg actggtttct ctgcaatggc gccaacccca	840
ggcttgccaa tactgcctat gtaaagggca agtgtgagaa	880
gctattctca tttcgctgac atacaggtag gactatgggg	920
gatgggacat ttgagtggga ctgagatagg aaaggcttga	960
aaagaaccca gaaacaccac caggaagttg gcaaagtaaa	1000
agaaaatgac ttccccctca aaggggcaatg agagggagag	1040
aaacaaacca aaatagaaga actagacttt ttagaaaatg	1080
agtattgcta	1090

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Year	1971	1972	1973	1974	1975	1976	1977	1978	1979	1980	1981	1982	1983	1984	1985	1986	1987	1988	1989	1990	1991	1992	1993	1994	1995	1996	1997	1998	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022	2023	2024	2025	2026	2027	2028	2029	2030	2031	2032	2033	2034	2035	2036	2037	2038	2039	2040	2041	2042	2043	2044	2045	2046	2047	2048	2049	2050	2051	2052	2053	2054	2055	2056	2057	2058	2059	2060	2061	2062	2063	2064	2065	2066	2067	2068	2069	2070	2071	2072	2073	2074	2075	2076	2077	2078	2079	2080	2081	2082	2083	2084	2085	2086	2087	2088	2089	2090	2091	2092	2093	2094	2095	2096	2097	2098	2099	2100
1971	1972	1973	1974	1975	1976	1977	1978	1979	1980	1981	1982	1983	1984	1985	1986	1987	1988	1989	1990	1991	1992	1993	1994	1995	1996	1997	1998	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022	2023	2024	2025	2026	2027	2028	2029	2030	2031	2032	2033	2034	2035	2036	2037	2038	2039	2040	2041	2042	2043	2044	2045	2046	2047	2048	2049	2050	2051	2052	2053	2054	2055	2056	2057	2058	2059	2060	2061	2062	2063	2064	2065	2066	2067	2068	2069	2070	2071	2072	2073	2074	2075	2076	2077	2078	2079	2080	2081	2082	2083	2084	2085	2086	2087	2088	2089	2090	2091	2092	2093	2094	2095	2096	2097	2098	2099	2100	

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hskcnq4  . . ILPSFRELALLFEHVQRARNGGLRPLEVRRAPVPDGA PSRYPPVATCHRPGSTSFPCPG
hskcnq5  . . KPHLKAL . . . . HT . . . . . CSPTK . . . . . KEQGEAS . . .
hskcnq2  TVTVPMYRLIPPLNQLELLRNLKSKSGLAFRKDPPPEPSP . . . . .
hskcnq3  VVSFPFFR . . . . . KEQLEAA . . . . .
hskcnq1  . . IRKAP . . RSHTLLS . . . . . PSPKPKKSUVVVKKKKFKLDKDNQV

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FIG. 5B

hskcnq4 ESSRMGIKDRIRMGSSQRRTPGS.KQQLAPPTMPTSPSSEQVGEATSPTKVQKSWSFNDR
hskcnq5 SSQKLSFKERVIRMASPRGQSIKS.RQ..ASVGDRRSPSTDITAEG.SPTKVQKSWSFNDR
hskcnq2 .SOKVSLKDRV.FSSPRGVAAKG.KGSPQAQTVRRSPSADQSLE.DSPSKVPKSWSFQDR
hskcnq3 SSQKLGLLDRVRLSNPRGSNTKG.KLF.....TPLNVDAIE.ESPSKEPKPVGLNNK
hskcnq1 PGEKMLTVPHITCDPPEERRLDHFSVDGYDSSVRKSET...LLEVSMPE.....

hskcnq4 TRFRASLRL....KPRTSABDA.PSEEVAAEKSYQCELTVDDEMPAVKTVIRSTRIRIKK
hskcnq5 TRFRPSLRLKSSQPKVIDADTALGTDDVYDEKGCQCDVSVEDLTPPLKTVIRATRIRIKK
hskcnq2 SRARQAFRIKGAASR.QNSEEASLPGEDIVDDKSCPCFVTEDETPGLKVSIRAVCMVRE
hskcnq3 ERFRTAFRMKAYAFWQ..SSEDAGTGDPMAEDRGYGNDFPIEDMIPTLKAAIRAVRILQK
hskcnq1 .HF...MRTNSFAEDLDLEGETLLT..PITH.....ISQREHHKATIKVIRRMQY

hskcnq4 LVAKRKFKETLRPYDVKDVIEWQYSAGHLDMGRIKSLQTRMDQIVGRG...PGDR.KARE
hskcnq5 HVAKRKFKETLRPYDVKDVIEWQYSAGHLDMGRIKSLQTRMDQILGKGQI.TSDK.KSRE
hskcnq2 LVSKRKFKESLRPYDVMVIEWQYSAGHLDMGRIKSLQSRMDQIVGRGPA.ITD..KDR.
hskcnq3 RLYKKKFKETLRPYDVKDVIEWQYSAGHLDMGRIKSLQTRMDMIFTPGPP.STPKHKKSQ
hskcnq1 FVAKKKFQOARKPYDVRDVIEWQYSAGHLNLMVRIKELORRLDQSIGKPSLFISVSEKSKD

hskcnq4 KGDKG.....PSDAEVVEISMMGRVVKVE..KQVQSIEHKLDLLLGFY
hskcnq5 ...KI.....TAEHETDLDLSMLGRVVKVE..KQVQSIESKLDCLLDIY
hskcnq2 ..TKG.....PAEAELPEDPSMMGRLGKVE..KQVLSMEKKLDLFLVNIY
hskcnq3 KGSFTFPSQQSPRNEPYVARPSTSEI.EDQSMGKFKVE..RQVQDMGKKLDLFLVDMH
hskcnq1 RGSNTIGARLNRVEDKVTQLDQRLALITD...MLHQLLSLHGGSTPGSGGPPREGGAHIT

hskcnq4 SRCLRSGT..SA.SLGAVQVPLFDPDITSYHSFVDEH..EDISVSAQTLS.ISRSVSTNM
hskcnq5 QQVLRKGS.SALALASFQIPPFCEQTSYQSPVDS..KDLGSAQNSGCLSRSTSANI
hskcnq2 MQ..RMGIP.PTETEAYFGAK..EPEAPPYHSPEDS..RE...HVDRHGCIVKIVRSSS
hskcnq3 MQHMER.....LQVQVTEYYPTKGTSSPAEAEKKEDNRYSDLKTIICNYSETGP
hskcnq1 QPCGSGGSVDPELFLPSNTLPTYE.QLTVPRRGPDIGS~~~~~

hskcnq4 D~~~~~
hskcnq5 SRGLQFI..LTPNEFSAQTFYALSPTMHSQATQVPISQSDGSAVAATNTIANQINTAPKP
hskcnq2 STGQKNF..SAPPAAPP...VQCPPSTSWQPQSHPRQGHGTSPVGDHGSVLRIPPPPAH
hskcnq3 PEPPYSFHQVTIDKVSPYGFFAHDPVNLPRGGPSS.GKVQATPPSSATTYVERPTVLPIL
hskcnq1 ~~~~~~

hskcnq4 ~~~~~~
hskcnq5 AAPTTLQIPPLPAIKHLRPETLHPNPAGLQESISDVTTCLVASKENVQVAQSNLTKDR
hskcnq2 ERSLSAYGGGNRASMEFLRQEDTPGCRPPEGTLRDSDTISIPSDHEELERSFSFGFSIS
hskcnq3 TLLDSRVSCHSQADLQG.PYSDRISPRQRRSITRSDTPLSLMSVNHEELERSPSFGFSIS
hskcnq1 ~~~~~~

hskcnq4 ~~~~~~
hskcnq5 SMRKSFDMGGETLLSVCPMVPK...DLGKSLSVQNLIRSTEELNIQLSGSESSGSRGSQ
hskcnq2 QSKENLDALNSCYAAVAPCAKVRPYIAEGESDTSDDLCTPCGPPRSATGEGPFQDVGWA
hskcnq3 QDRDDYVFGPN...GGSSWMREKRYLAEGETDTDTPFTPSGSMPLSSTGDGIDS SVWTP
hskcnq1 ~~~~~~

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FIG. 5C

hskcnq4 ~~~~~
hskcnq5 DFYPKWRESKLFITDEEVGPEETETDTFDAAPQPAREAAAFASDSLRTGRSRSSQSICKAG
hskcnq2 GPRK~~~~~
hskcnq3 SNKPI~~~~~
hskcnq1 ~~~~~

hskcnq4 ~~~~~
hskcnq5 ESTDALSLPHVKLK
hskcnq2 ~~~~~
hskcnq3 ~~~~~
hskcnq1 ~~~~~

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"02099860"

FIG. 6A

Human RNA Master Blot

TABLE 1

	1	2	3	4	5	6	7	8
A	whole brain	amygdala	Caudate nucleus	cerebellum	cerebral cortex	frontal lobe	hippocampus	medulla oblongata
B	occipital pole	putamen	substantia nigra	temporal lobe	thalamus	Subthalamic nucleus	spinal cord	
C	heart	aorta	Skeletal muscle	colon	bladder	uterus	prostate	stomach
D	testis	ovary	pancreas	pituitary gland	adrenal gland	thyroid gland	salivary gland	mammary gland
E	kidney	liver	small intestine	spleen	thymus	peripheral leukocyte	lymph node	bone marrow
F	Appendix	lung	trachea	placenta				
G	fetal brain	fetal heart	fetal kidney	fetal liver	fetal spleen	fetal thymus	fetal lung	
H	yeast total RNA	yeast tRNA	<i>E. coli</i> rRNA	<i>E. coli</i> DNA	Poly r(A)	human C ₀ t DNA	human DNA	human DNA

T04250-02099850

FIG. 6B

	1	2	3	4	5	6	7	8
A	•		•		•	•	•	
B	•	•		•				
C			•					
D								
E								
F								
G								
H				•				

T04250" 02099860

099860 02099860

FIG. 7

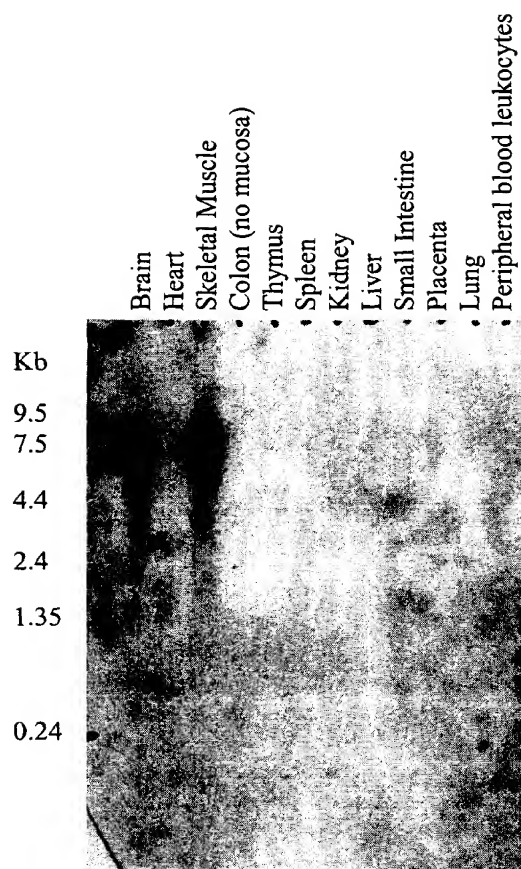
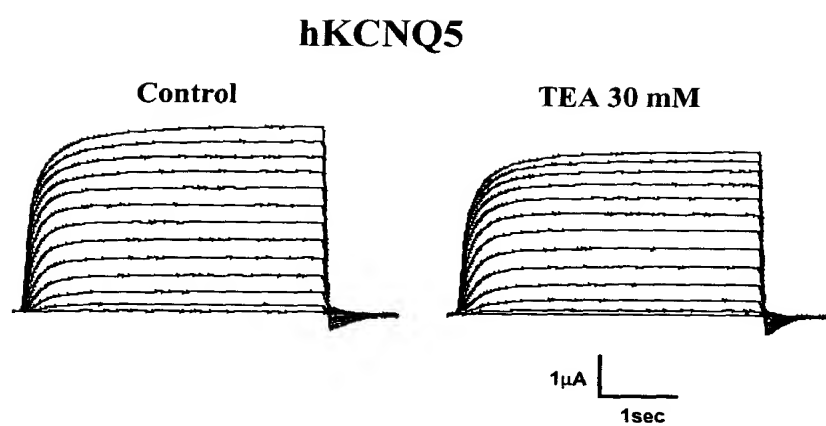


FIG. 8



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Antisense



FIG. 9A



FIG. 9B



FIG. 9C

Sense



FIG. 9D

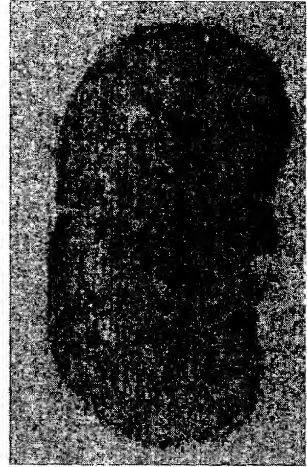


FIG. 9E



FIG. 9F

FIG. 10A

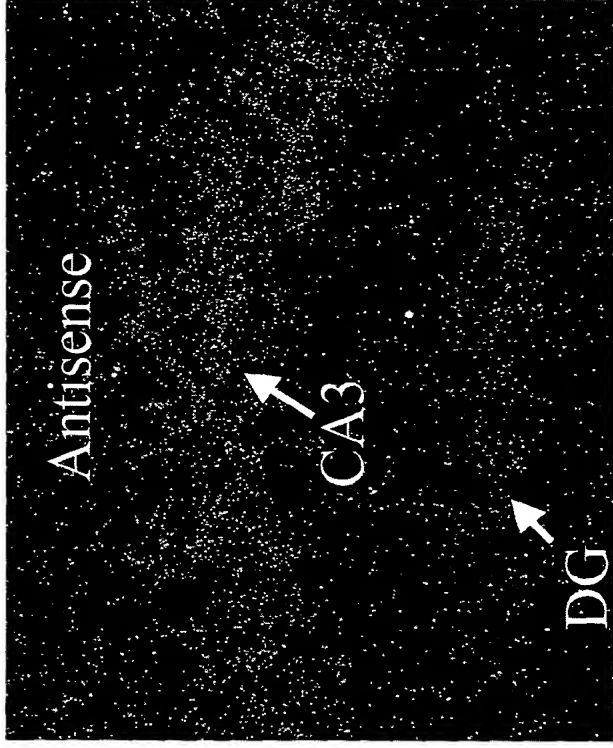


FIG. 10B

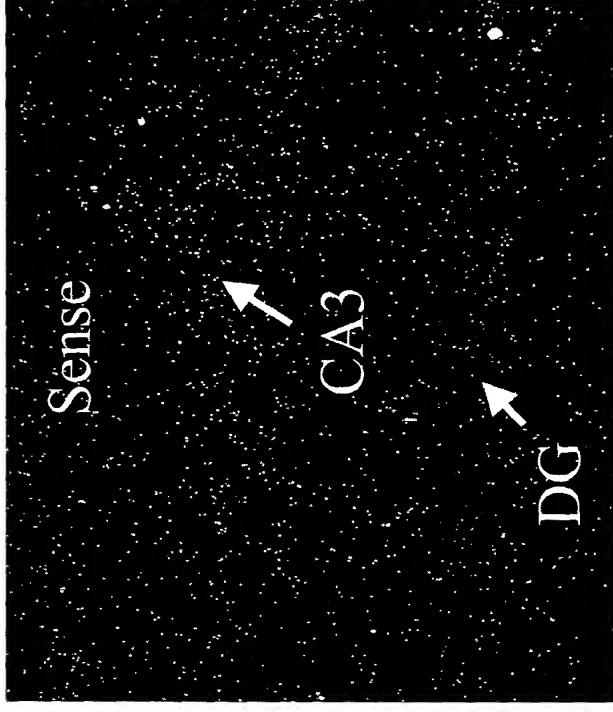


FIG. 11

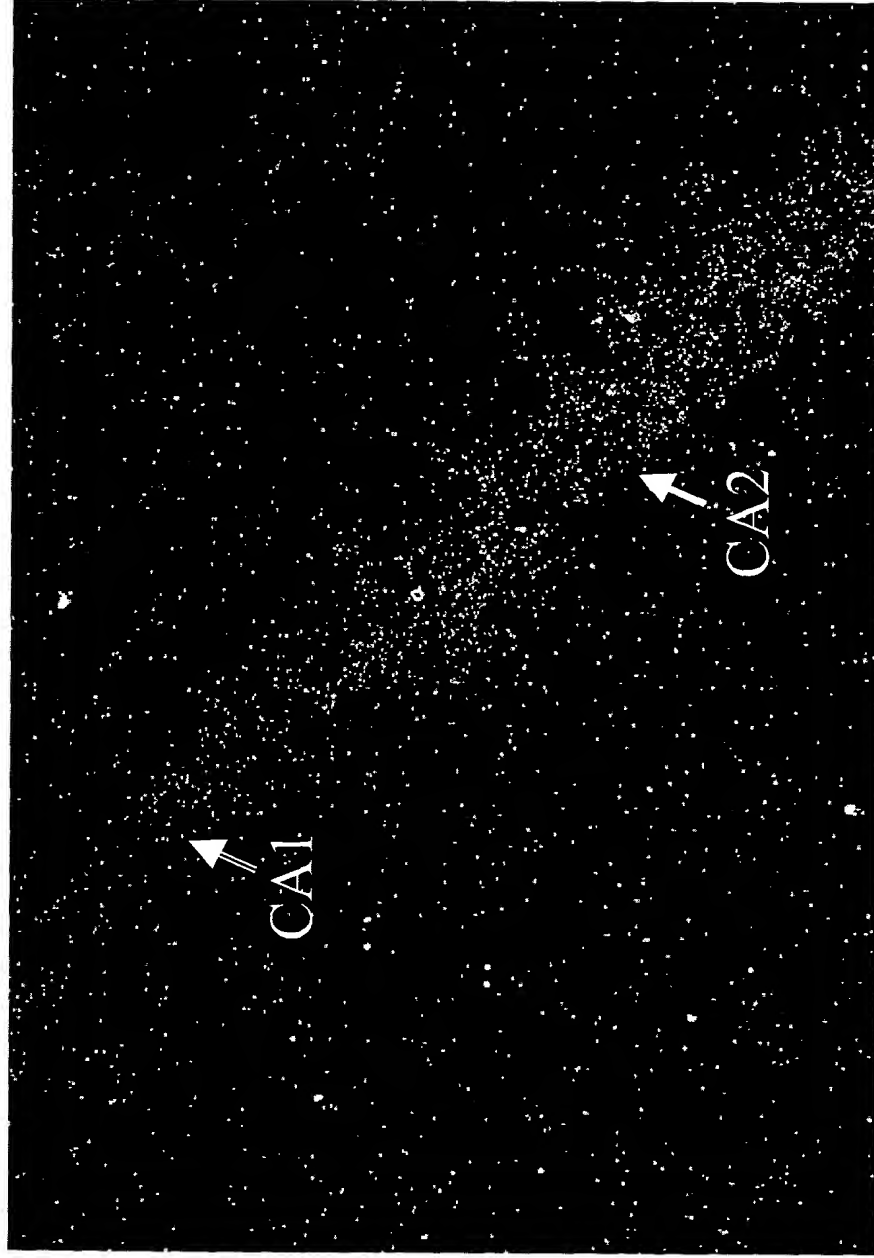


FIG. 12A

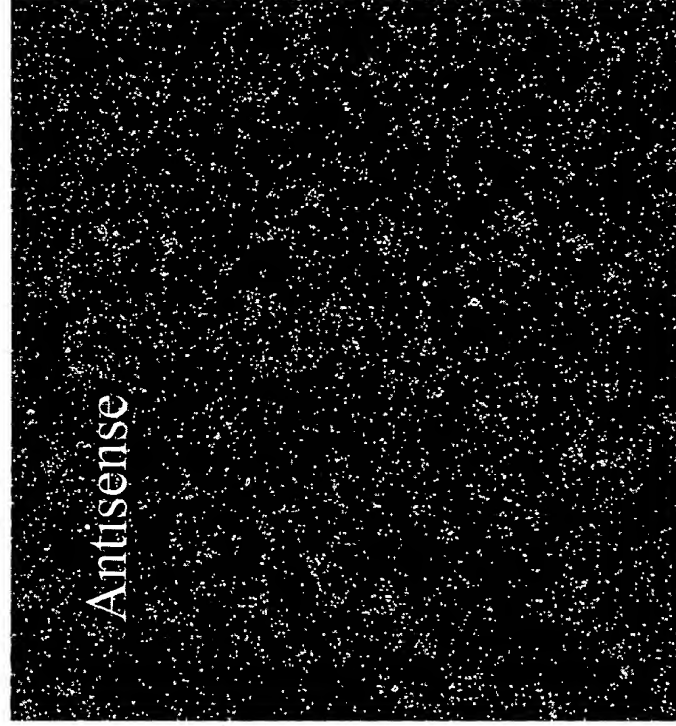


FIG. 12B

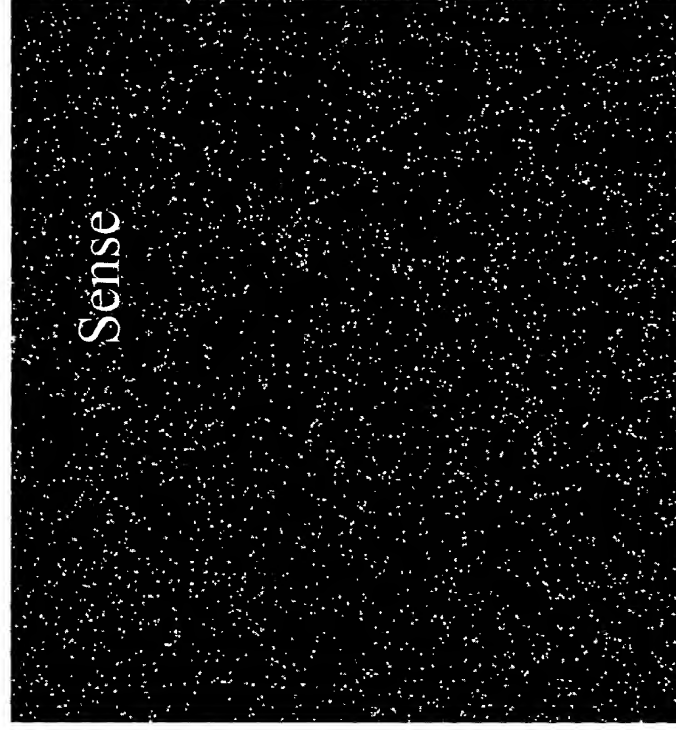


FIG. 13A

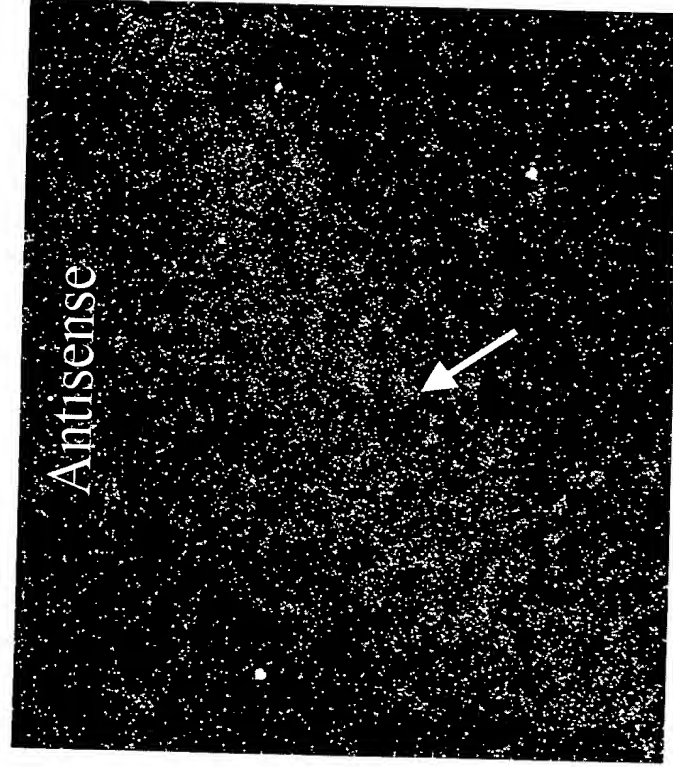


FIG. 13B

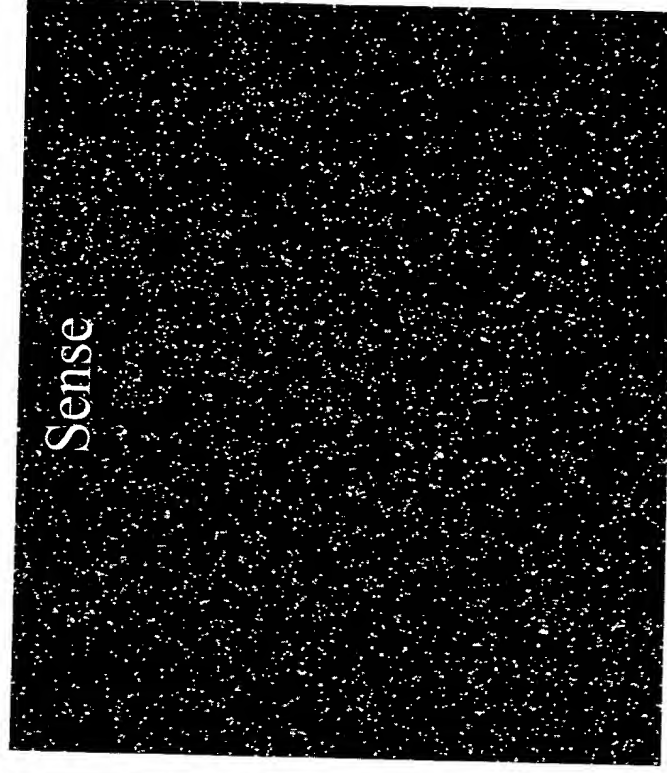


FIG. 14A

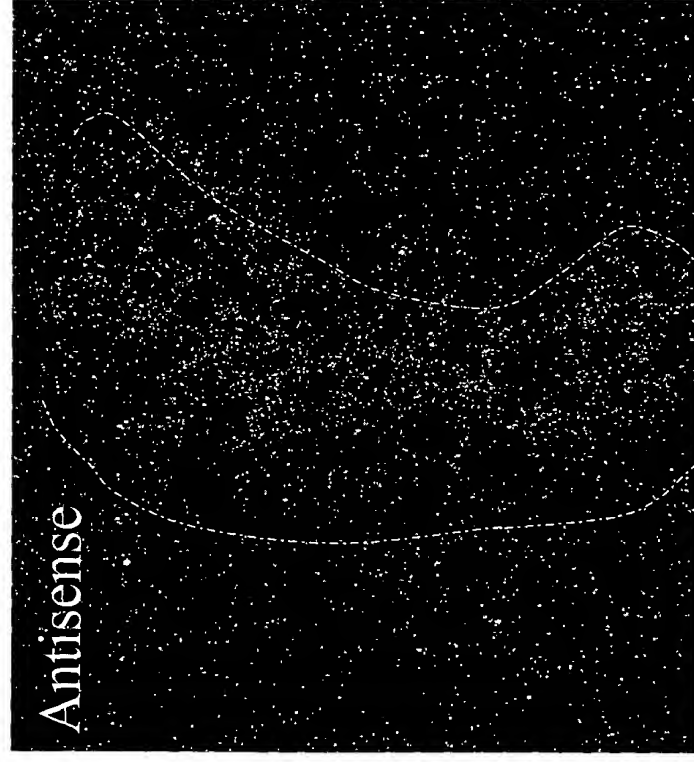


FIG. 14B

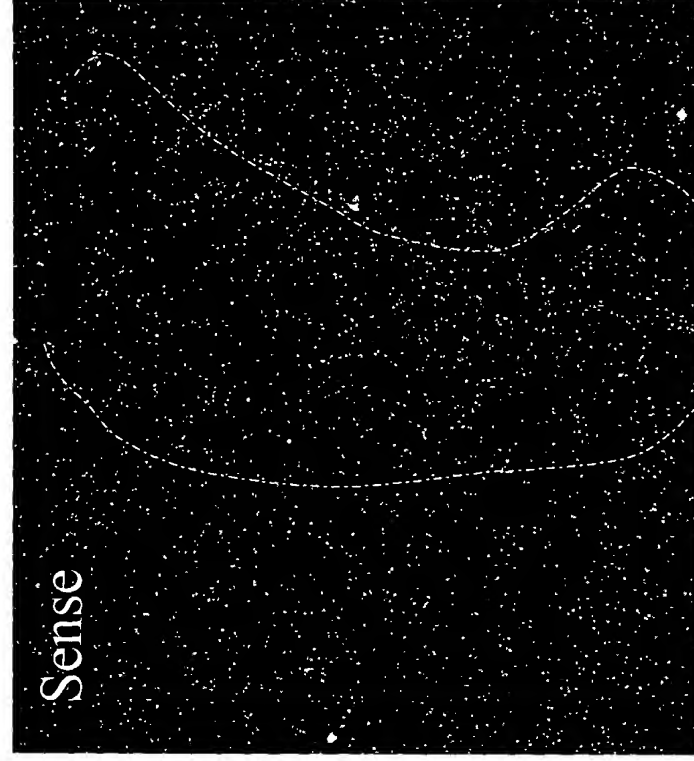


FIG. 15A

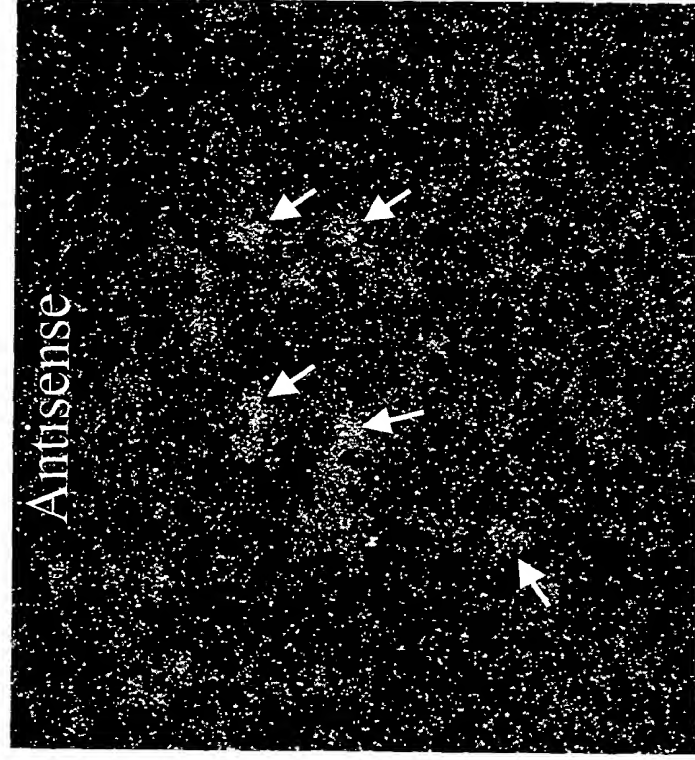


FIG. 15B

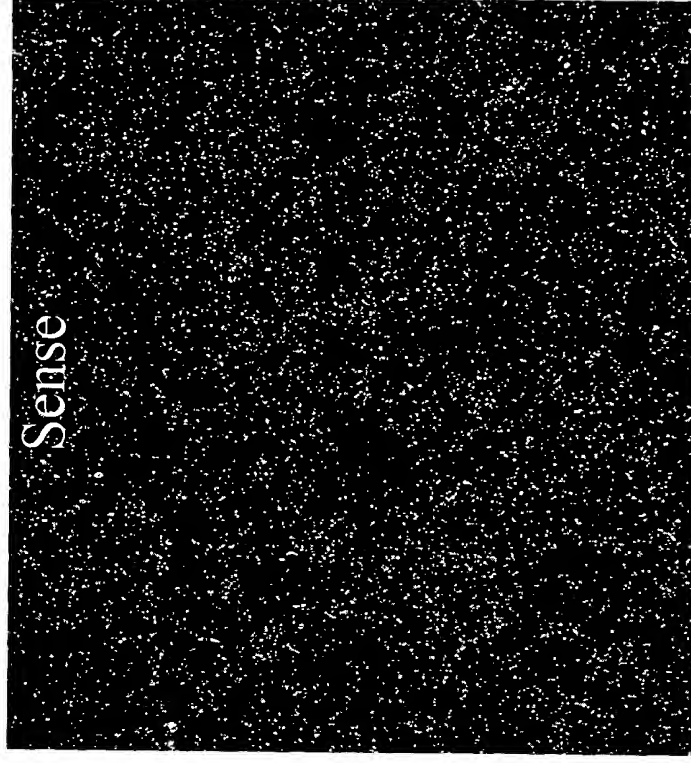


FIG. 16A

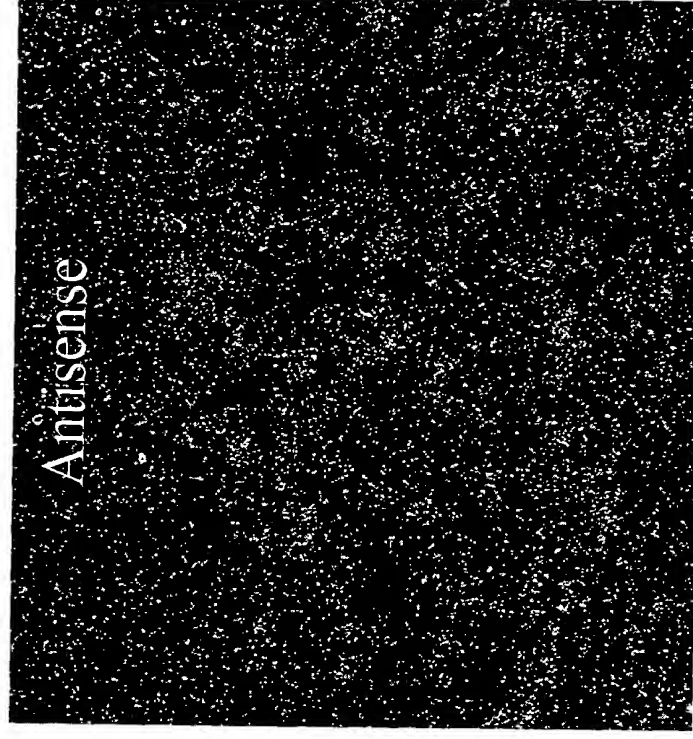


FIG. 16B

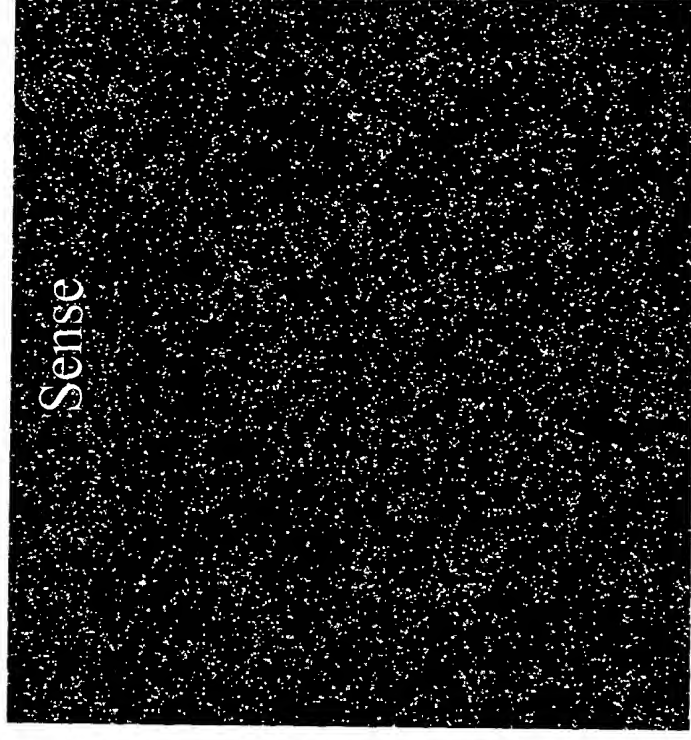


FIG. 17A

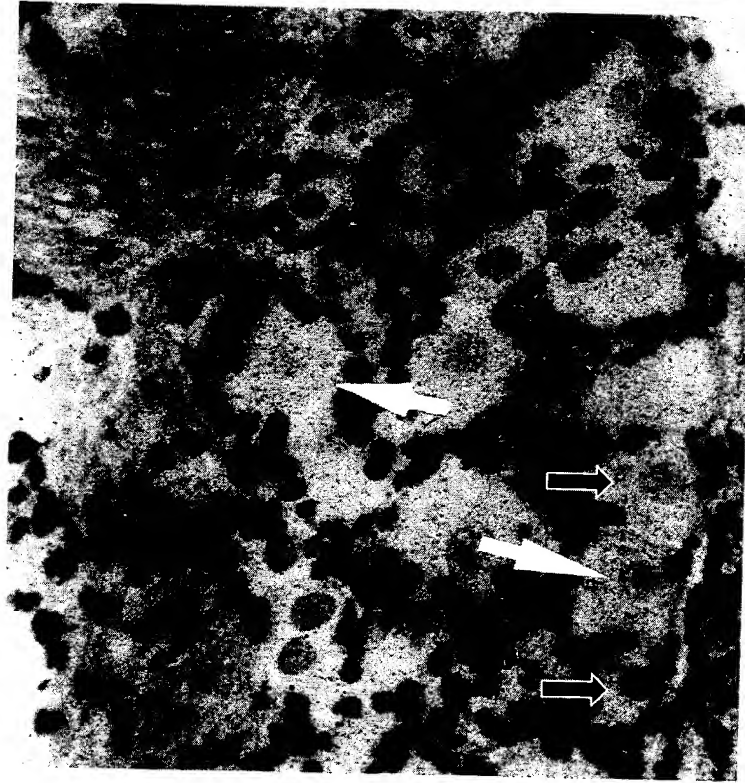


FIG. 17B

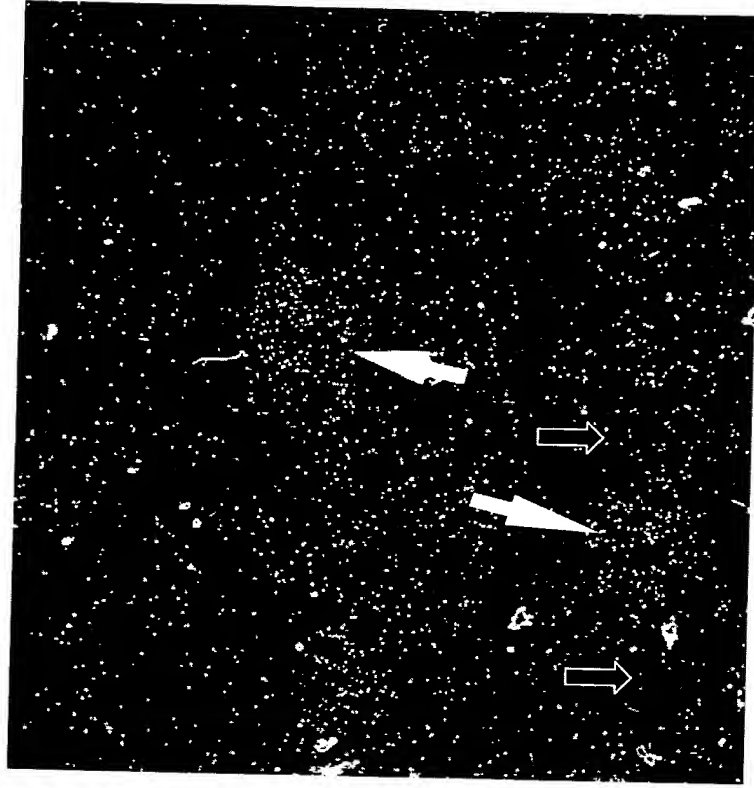


FIG. 18A

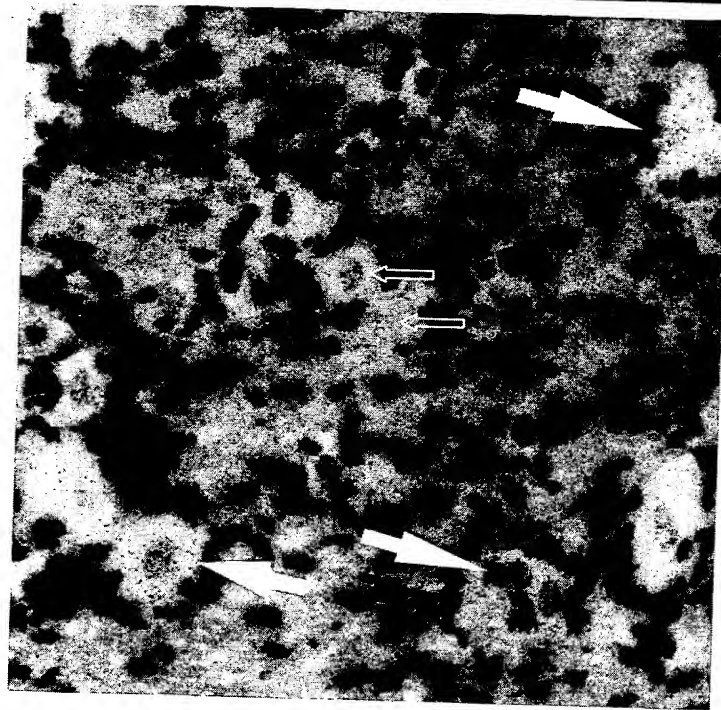


FIG. 18B

